

09458366-120999

1 GGCACCAGGAGATCTAGCTCAAATTAATGTTGCCCCCTAGCTGCTAAAGGACAGAGACCCCTCAGACTGATGAAATGCGCTCAGAATTACTTAGACAA  
97 AGCGGATATTTCCTACTCTCTTCCCTTTTCTCTGTTTTTCTAGTGAAGAGACCTGAAAGAAAAAGTAGGGAGAACATAATGAGAACAAATACG  
193 GTAATCTCTTCATTTCCTAGTTCAAGTCTGGACTTGGGACTTAGGAGGGGCAATGGAGCCGCTTAGTGCCTACATCTGACTTGGACTGAAATATA  
289 CGTGAGAGACAAGATTGTCTCATATCCGGGAAATCATACCTATGACTAGGACGGGAAGAGCAAGCACTGCCTTTACTTCACTGGGAATCTCGGC  
385 CTCAGCCTGCAAGCCAAAGTGTTTCAAGTGAAGAAAAGCAAGAGAATAAGCTAATACTCTGTCTCTGAACAAGGCAGCGGCTCCTTGCTAAAGCTACT  
481 CCTTGATCGATCCTTTGCACCGGATTGTTCAAAGTGGACCCAGGGGAGAAAGTCGGAGCAAAGAACTTACCACCAAGCAGTCCAAGAGGCCAGAA  
577 GCAAACTGGAGGTGAGACCCAAAGAAAGCTGGAACCATGCTGACTTTGTACACTGTGAGGACACAGAGTCTGTTCTGGAAAGCCAGTGTCAAC  
L E V R P K E S W N H A D F V H C E D T E S V P G K P S V N 30  
673 GCAGATCAGGAAGTCGGAGCTCCCAAATCTGCCGTGTATGTGGGACAAAGGCCACTGGCTATCACTTCAATGTGATGACATGTGAAGGATGCAAG  
A D E E V G G P Q I C R V C G D K A T G Y H F N V M T C Z G C K 62  
769 GGCCTTTTCAGGAGGGCCAACAAAGCAAGCCCGGCTGAGGTGCCCTTCCGGAAGGGCGCTGCGAGATCACCCGGAAGACCCGGGACAGTGC  
G F F R R A M K R N A R L R C F F R K G A C E I T R K T R R Q C 94  
865 CAGGCCTCCCGCTCGCAAGTGCCTGGAGAGCGGCATGAAGAAGGAGATGATGTCCGACGAGGCGCTGGAGGAGAGCGGGCTTGATCAAG  
Q A C R L R K C L E S G M K K E H I H S D E A V E E R R A L I K 126  
961 CGGAAGAAAAGTCAACCGACAGGGACTCAGCCACTGGGAGTGCAGGGGTGACAGAGCAGCGGATGATGATCAGGGAGCTGATGGACGCTCAG  
R K K S E R T G T Q P L G V Q G L T E E Q R M H I R E L M D A Q 158  
1057 ATGAAAACCTTTGACACTACCTTCTCCCATTTCAAGAAATTCGGCTGCCAGGGGTGCTTAGCACTGGCTCGGAGTGGCAGAGCCTCTGCAGGCC  
M K T F D T T F S H F K N F R L P G V L S S G C E L P E P L Q A 190  
1153 CCATCGAGGGAAGAAGCTGCCAAGTGGAGCCAGGTCCGGAAGATCTCTGCTCTTTGAAGGTCTCTCTGCAAGCTCGCGGGGAGGATGGCACTGT  
P S R E E A A K W S Q V R K D L C S L K V S L Q A A G G G W Q C 222  
1249 CTGGAAGTACAAACNCCAGCCGACAGTGGCGGAAGAGATCTTCTCCCTGCTGCCCCACATGGCTGACATGTCAACCTACATGTTCAAAGGCATC  
L E L Q T P S R Q W R K E I F S L L P H M A D M S T Y M F K G I 254  
1345 ATCAGCTTTGCCAAAGTCATCTCTACTTCAGGGACTTGCCTATCGAGGACCAGATCTCCCTGCTGAAGGGGGCGCTTTCGAGCTGTGTCAACTG  
I S P A K V I S Y F R D L P I E D Q I S L L K G A A F E L C Q L 286  
1441 AGATTCAACACAGTGTTCACCGCGGAGACTGGAACCTGGGAGTGTGGCCGGCTGTCTACTGCTTGGGAAGACACTGCAGGTGGCTTCCAGCACTT  
R F N T V F N A E T G T W E C G R L S Y C L E D T A G G F Q Q L 318  
1537 CTACTGGAGCCCATGCTGAAATCCACTACATGCTGAAGAAGCTGCAGCTGCATGAGGAGGAGTATGTGCTGATGCAGGCCATCTCCCTCTTCTCC  
L L E P H L K F H Y M L K K L Q L H E E E Y V L M Q A I S L F S 350  
1633 CCAGACCGCCAGGTCTGCTGCAGCACCGGCTGGTGGACCAGCTGCAGGAGCAATTCGCCATTACTCTGAAGTCTACATTGAATGCAATCGGCC  
P D R P G V L Q H R V V D Q L Q E Q P A I T L K S Y I E C N R F 382  
1729 CAGCCTGCTCATAGGTTCTGTTCCTGAAGATCATGGCTATGCTCACCGAGCTCCGAGCATCAATGCTCAGCACACCCAGCGGCTGCTGGGCATC  
Q P A H R F L F L K I H A M L T E L R S I N A Q H T Q R L L R I 414  
1825 CAGGACATACACCCCTTTGCTACGCCCTCATGCAGGAGTGTTCGGCATCACAGGTAGCTGAGCGGCTGCCTTGGGTGACACCTTCGAGAGGCAG  
Q D I H P F A T P L M Q E L F G I T G S 434  
1921 CCAGACCCAGAGCCCTCTGAGCCGGCACTCCCGGGCAAGACAGATGGACACTGCCAAGAGCCGACAAATGCCCTGCTGGCTGTCTCCCTAGGGAA  
2017 TTCCTGCTATGACAGCTGGCTAGCATTCCTCAGGAAGGACATGGGGTGCCCC 2068

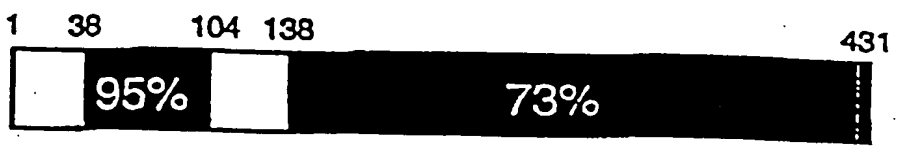
FIG. 1A

0945366.120999

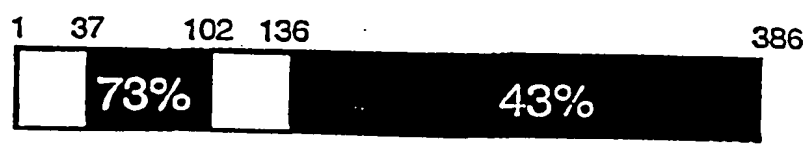
hSXR



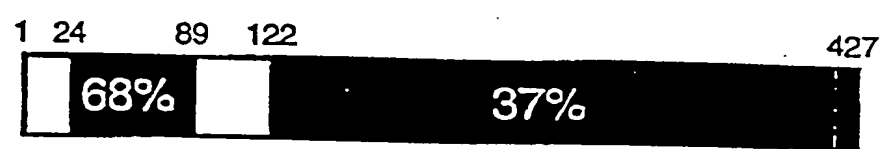
mPXR.1



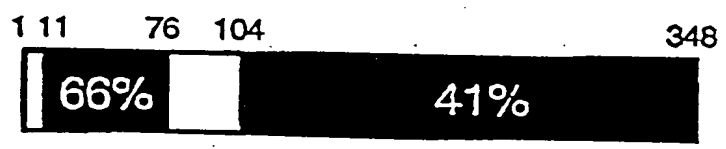
xBXR



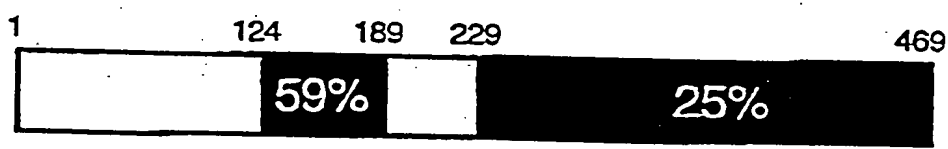
hVDR



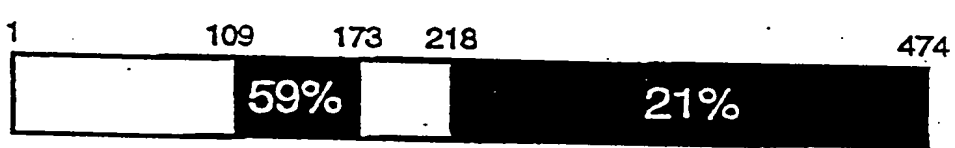
hCAR $\alpha$



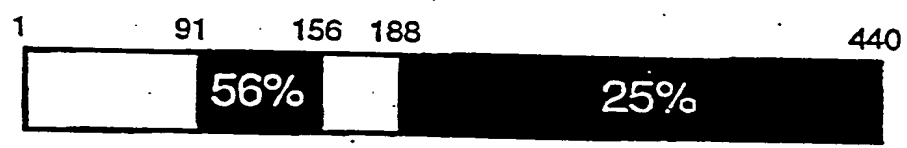
rFXR



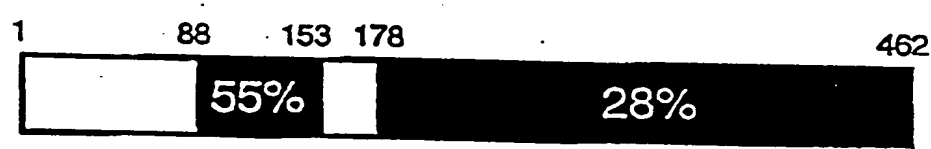
mPPAR $\alpha$



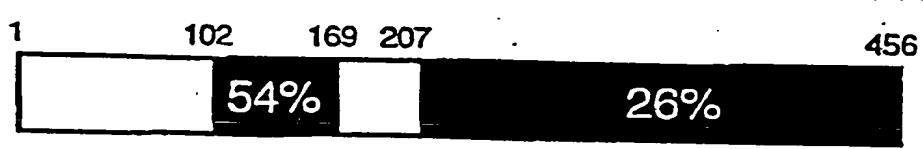
hLXR $\alpha$



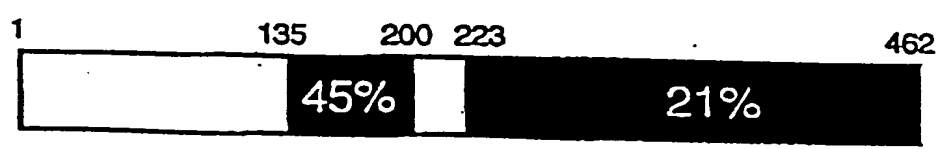
hRAR $\alpha$ 1



hTR $\beta$



hRXR $\alpha$



hGR $\alpha$

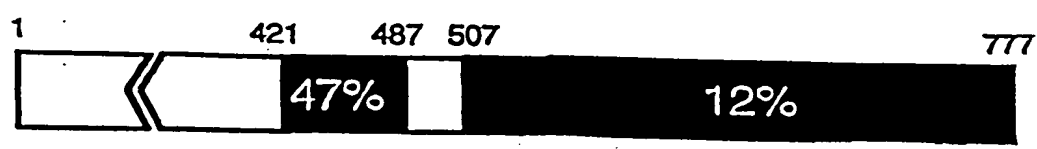


FIG. 1B

09452366-120999

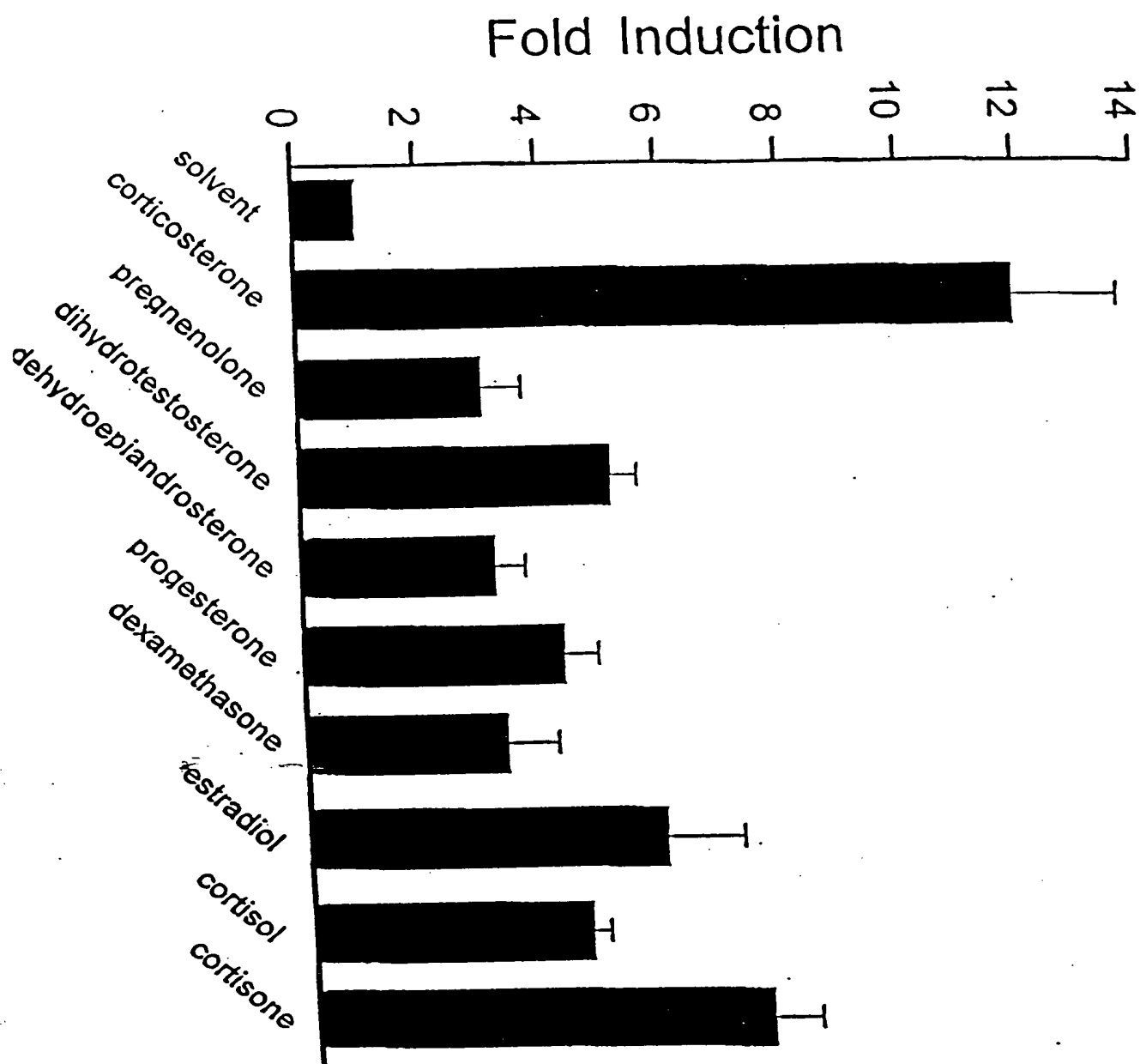


FIG. 2

# Fold Induction

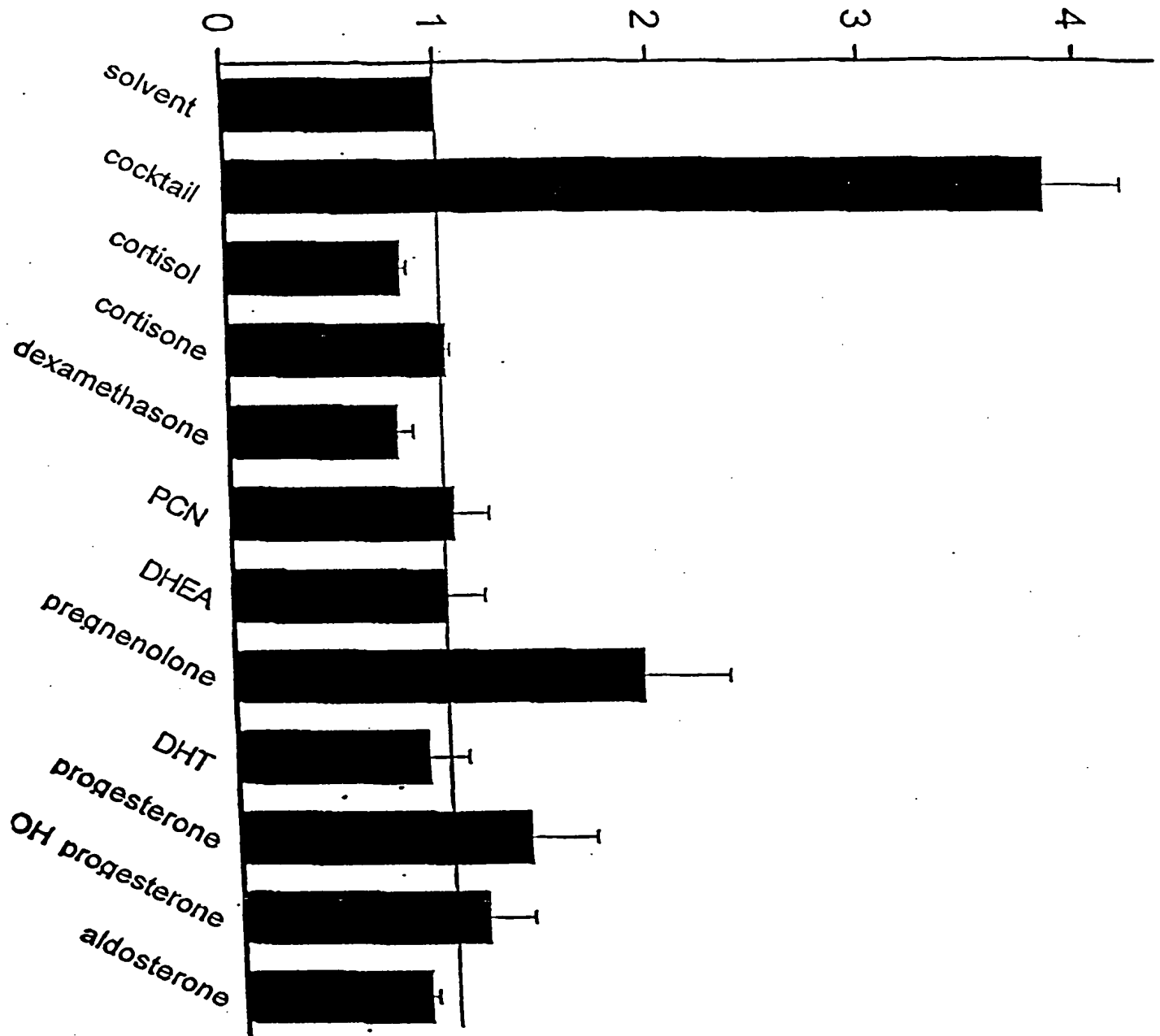


FIG. 3

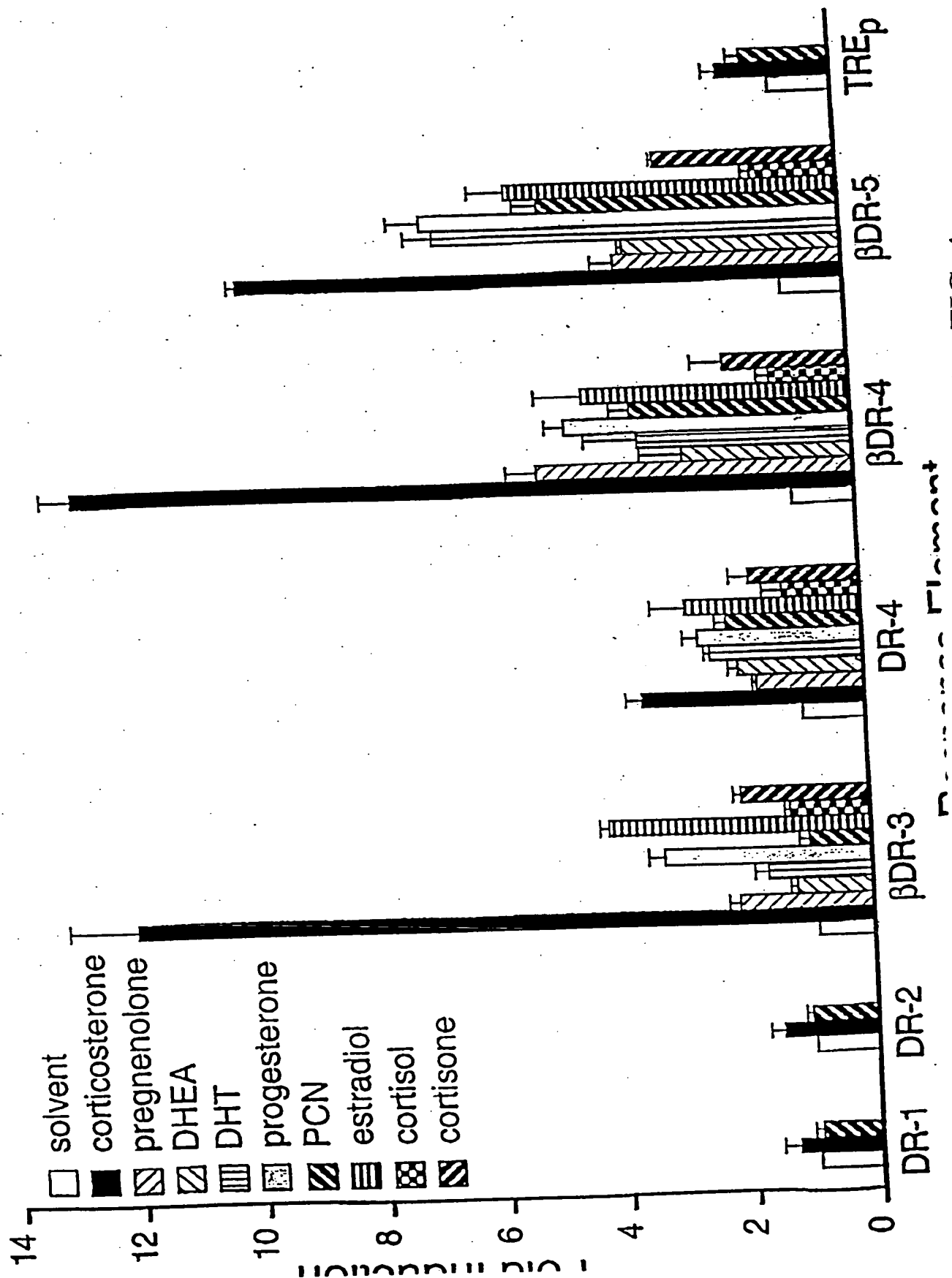


FIG. 4

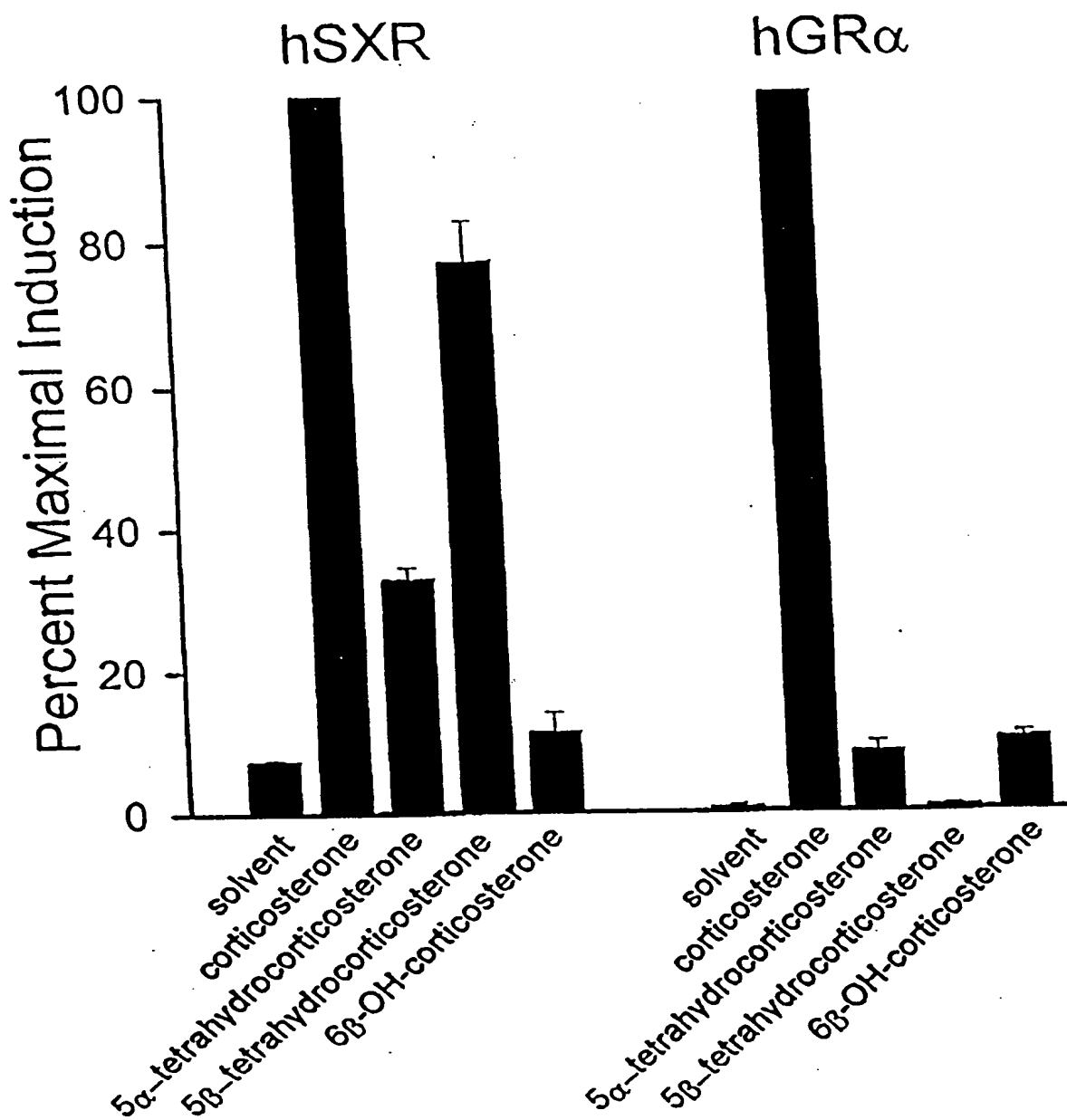


FIG. 5

DR-3

rCYP3A1  
rCYP3A2  
rUGT1A6

tagac AGTTCA tga AGTTCA tctac  
taagc AGTTCA taa AGTTCA tctac  
actgt AGTTCA taa AGTTCA catgg

DR-4

rbCYP2C1  
rP450R

caatc AGTTCA acag GGTTCa ccaat  
cac AGGTGA gctg AGGCCA gcagc AGGTCC aaa

DR-5

rCYP2A1  
rCYP2A2  
rCYP2C6  
hCYP2E1

gtgca GGTTCa actgg AGGTCA acatg  
gtgct GGTTCa actgg AGGTCA gtatg  
agtct AGTTCA gtggg GGTTCa gtctt  
gagat GGTTCa aggaa GGGTCA ttaac

FIG. 6A

CYP3A4  
CYP3A5  
CYP3A7

tagaata TGAACt caaagg AGGTCA gtgagtgg  
tagaata TGAACt caaagg AGGTAA gcaaaggg  
tagaata TTAACt caatgg AGGC.A gtgagtgg

FIG. 6B

0945366-12099

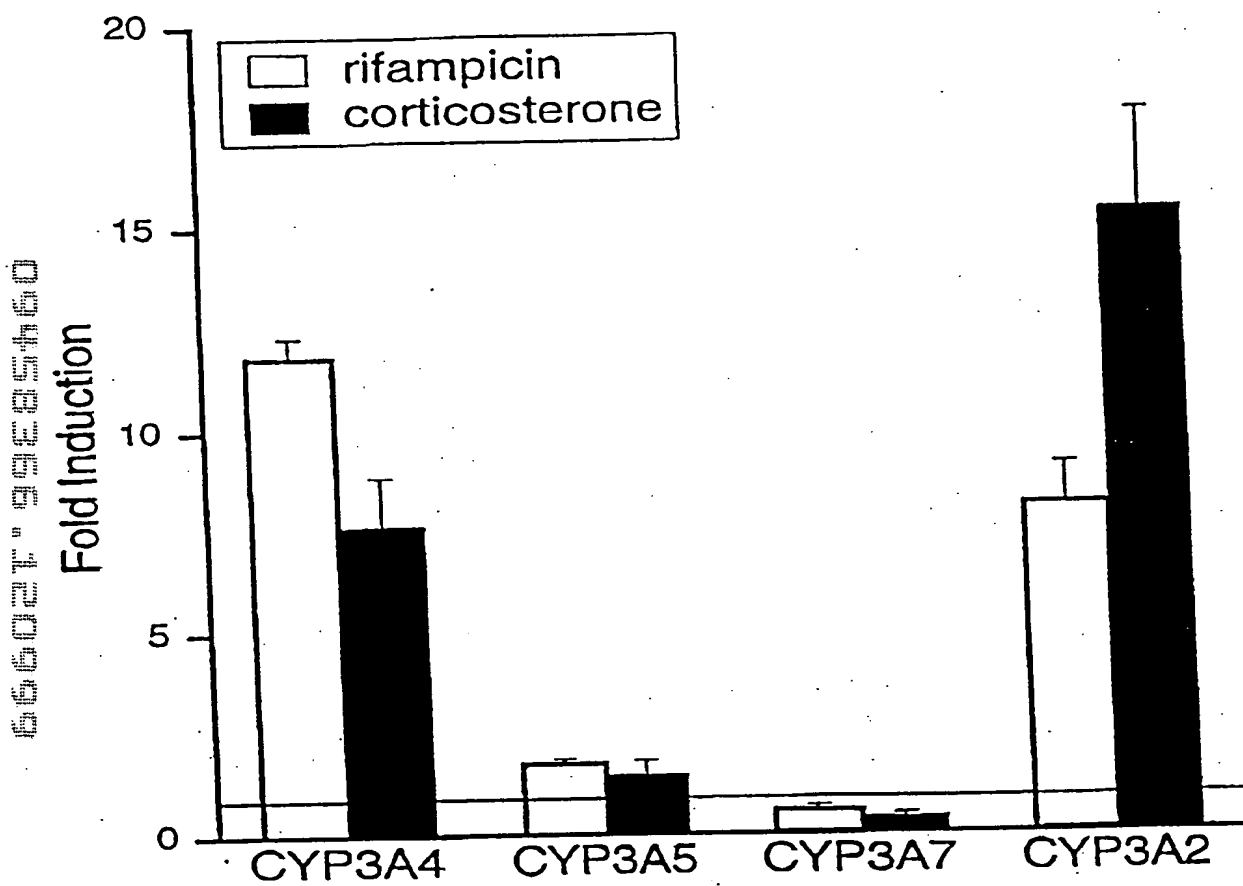
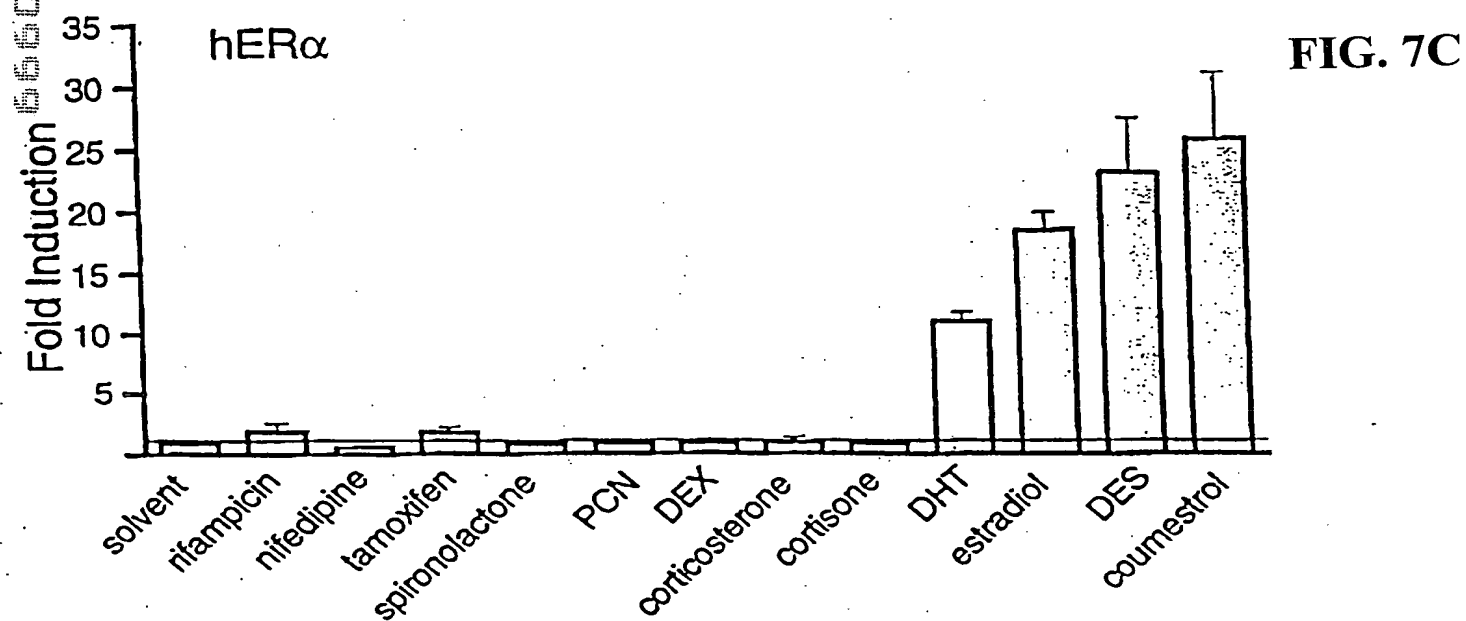
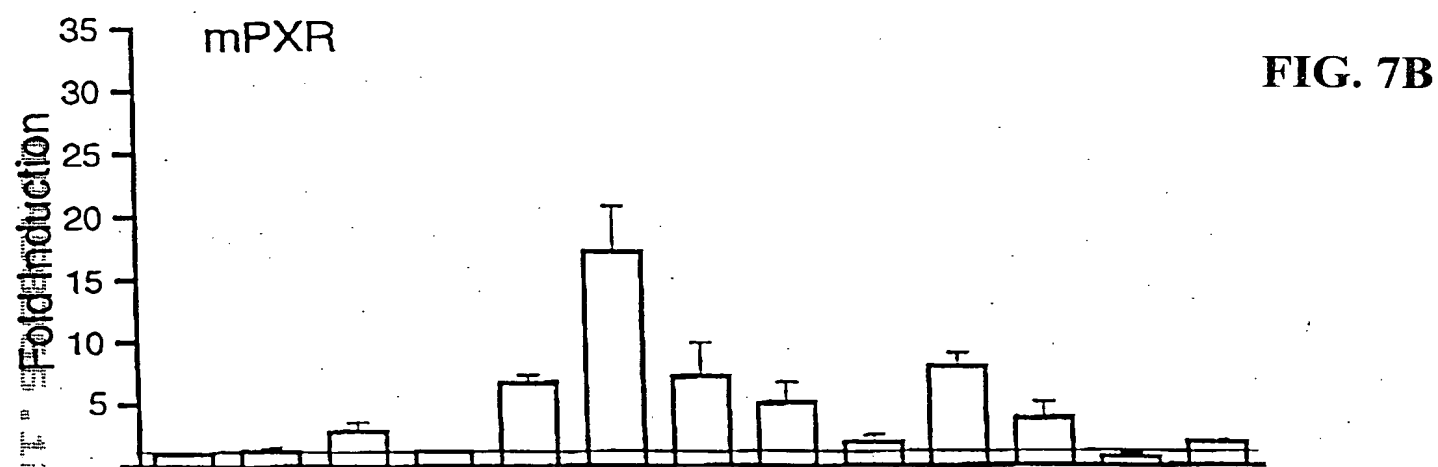
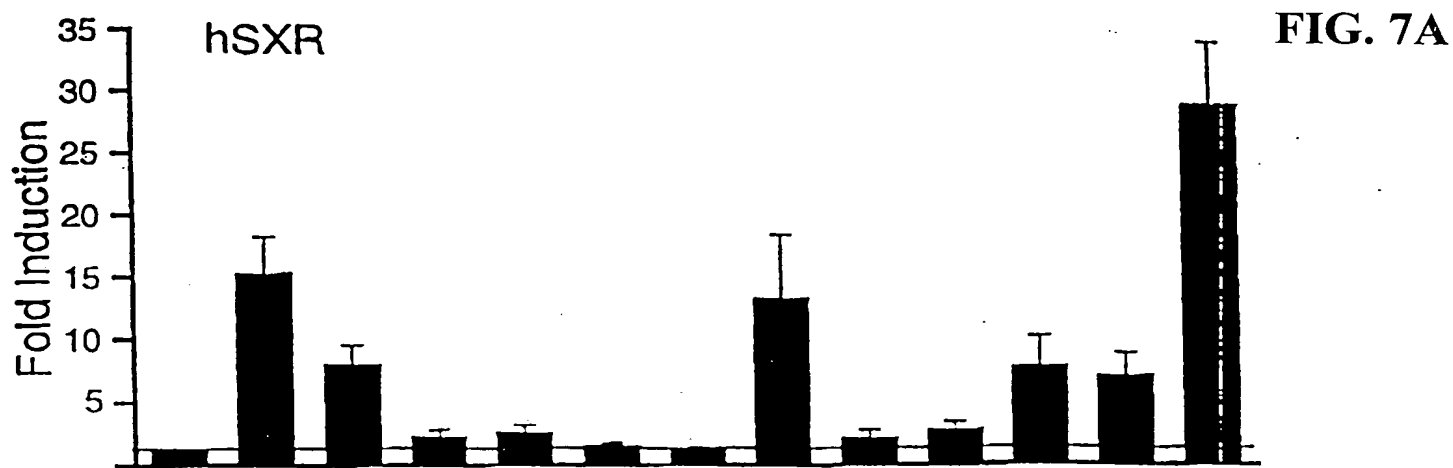


FIG. 6C





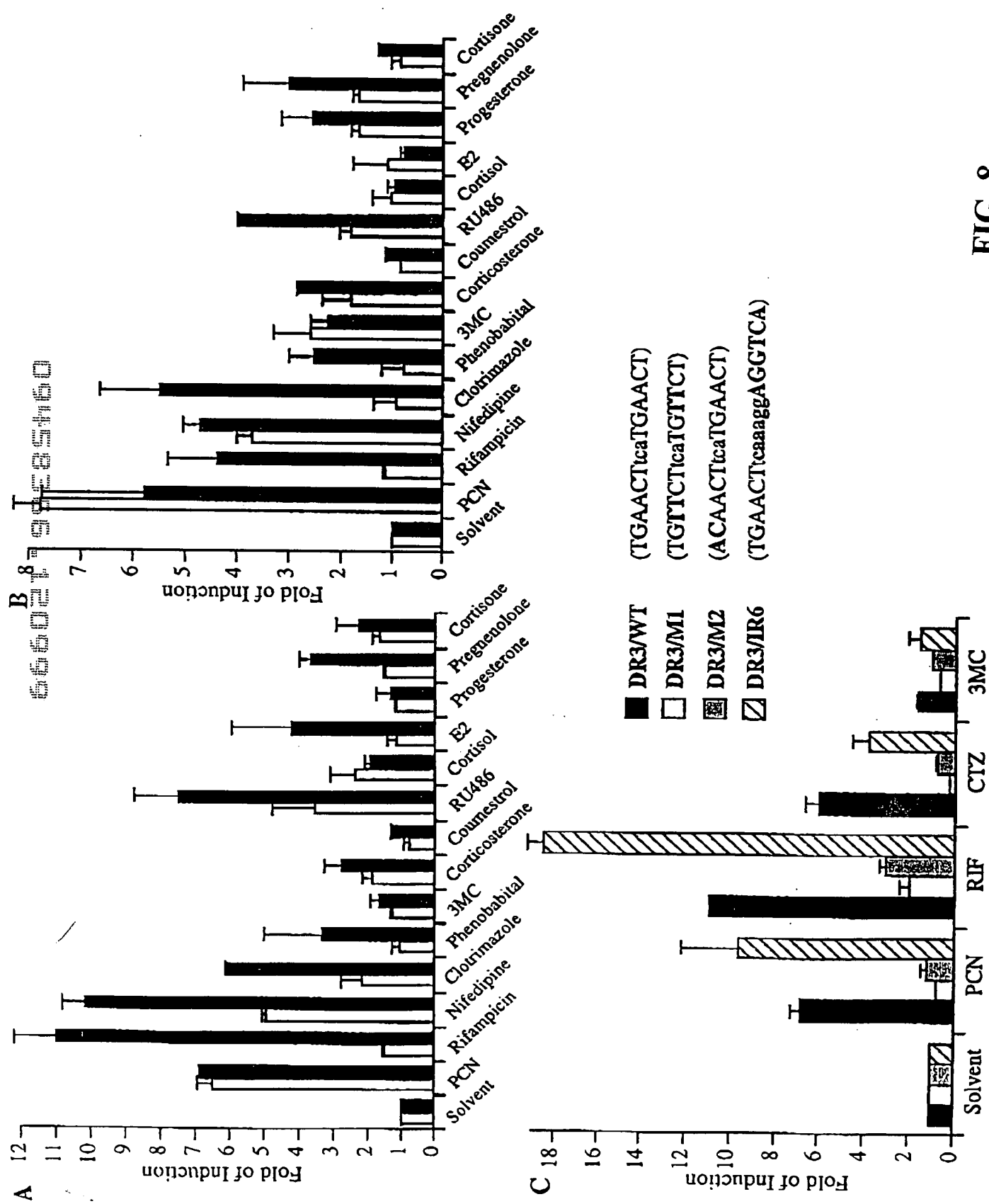


FIG. 8

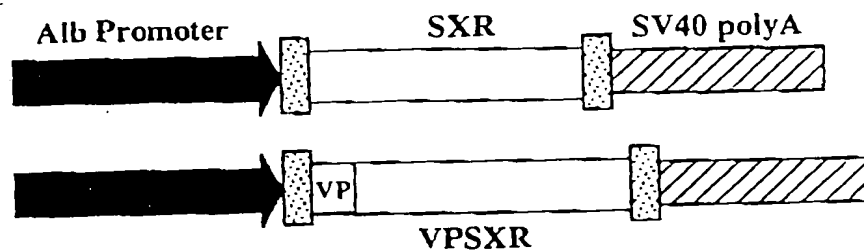


FIG. 9

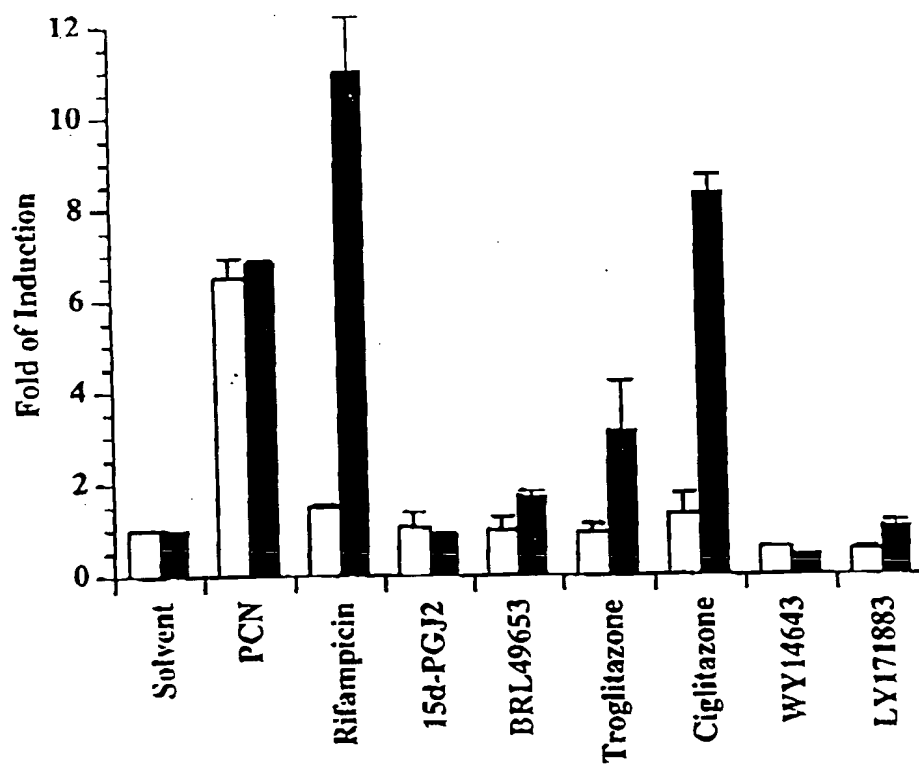
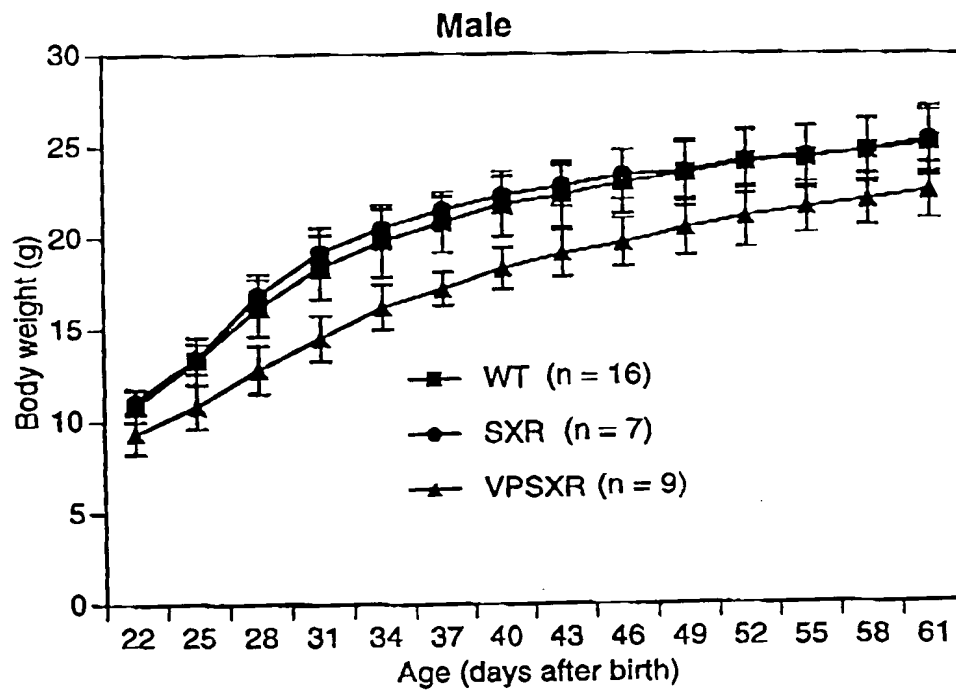
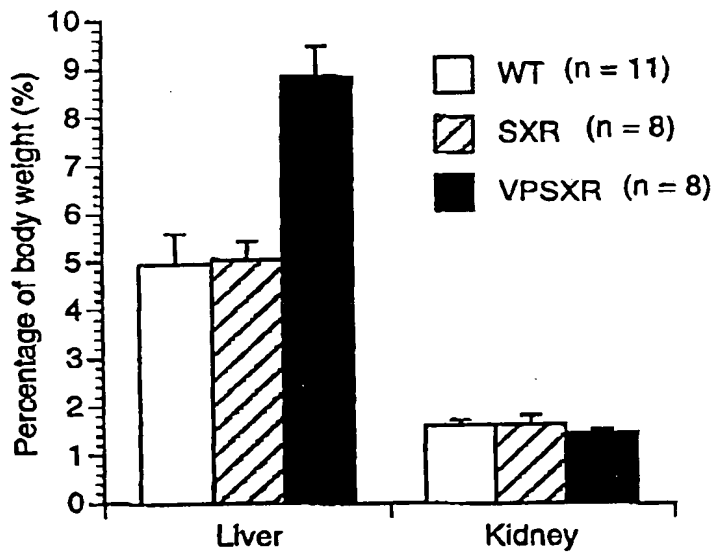


FIG. 10



**FIG. 11**



**FIG. 12**